This Page Is Inserted by IFW Operations and is not a part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

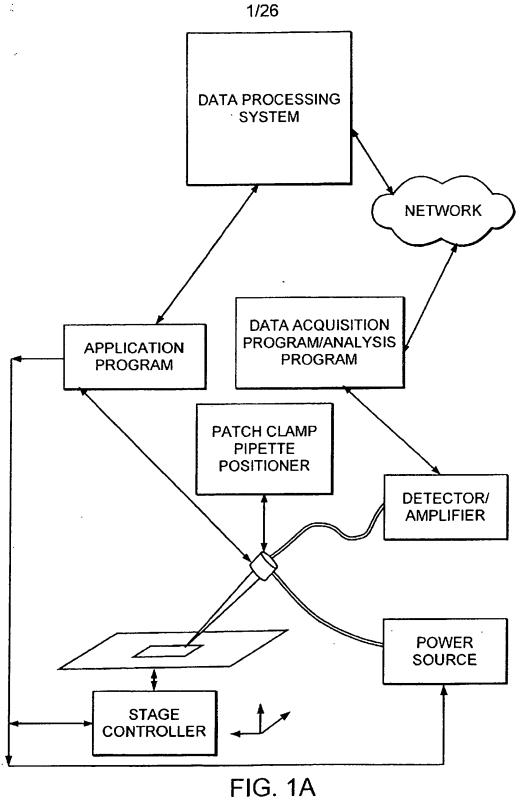
Defects in the images may include (but are not limited to):

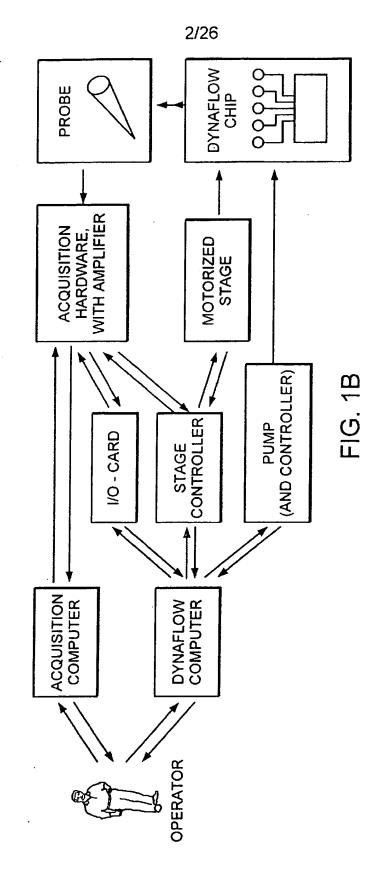
- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

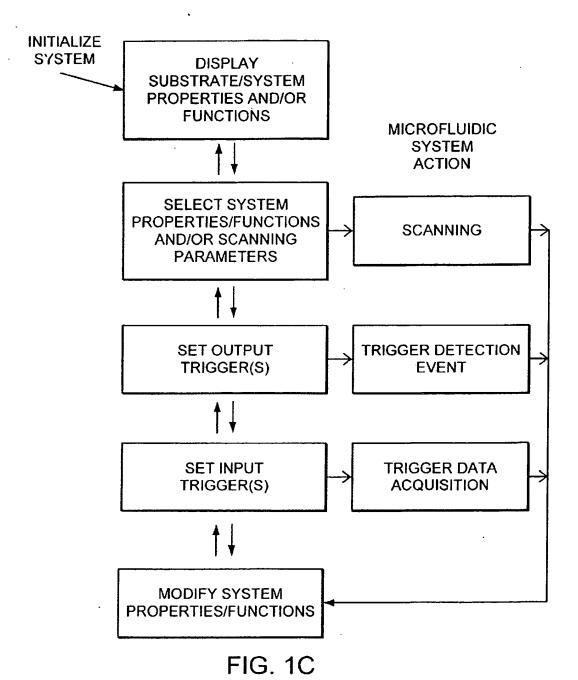
IMAGES ARE BEST AVAILABLE COPY.

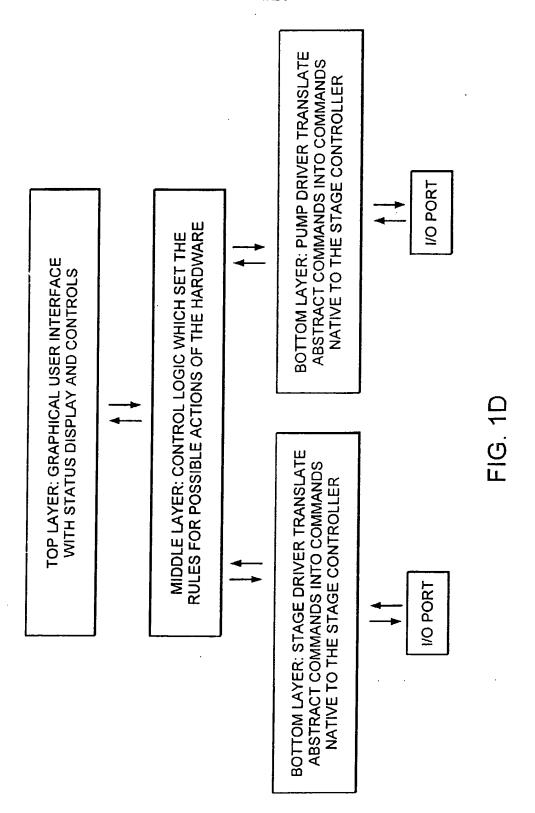
As rescanning documents will not correct images, please do not report the images to the Image Problem Mailbox.











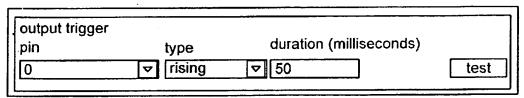
p DynaMan 0.5 m		
chip type setting	stage settings	
90µm.32 channels ▽	load edit	save

FIG. 2

Stage Settings		\boxtimes
main settings stage type po Virtual test stage C	rt maximum speed	(microns / second)

Setting stage, port and max speed.

FIG. 3



Output trigger settings.

FIG. 4

input trigger			
pin	type		
0		▽	test

FIG. 5

stage s	status	s					-	-																					
channe	el: 4	.	st	at	us	3 :	0)	ŗ	oro	gr	es	s:	()	5	sta	ge	e p	ro	g	re:	SS		1.	00	00	00	0
	11	ī	1	ī	-	1	ī	1	ŀ	ı	1	1	1	ī	1	1	ī	1	ī	1	ī	ı	ī	1	1		ī	-]
0,0									р	os	itic	on:	;	51	6,	51								4	10	00),4	00)

FIG. 6A

stage control		
set refpos A	set refpos B	
channel 2 ▽	⇒start	[][] stop
scan mode	time/chanr	nel
	∀	▽
output start trig	gger 🔲 output o	channel trigger
input start trigg	ger output o	channel tags

FIG. 6B

翻 DynaMan 0.5	<u>₹</u>
chip type setting stage settings 90μm.32 channels load edit save	
stage status	
channel: 1 status: 0 progress: 0 stage progress: 0.000000	
stage control]
set refpos B set refpos B	
channel 2	
scan mode time/channel continuous movement □ [6,450000 (6,450000)] □ □ □	
output start trigger output channel trigger input start trigger output channel tags	
Stage creation and connection ok.	

FIG. 7

Stage Settings			X
main settings stage type Virtual test stage ▽	port	maximum speed (micror	ns / second) test
output trigger type rising	pin 0	duration (milliseconds) ▼ 50	test
input trigger type rising ▽	pin 0	∀	test
pump settings pump type	port	▼	test
		ОК	Cancel

FIG. 8

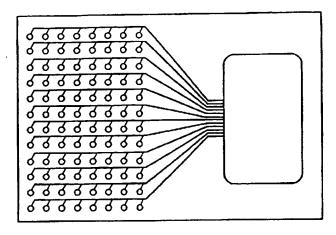


FIG. 9A

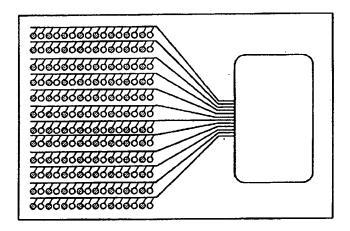


FIG. 9B

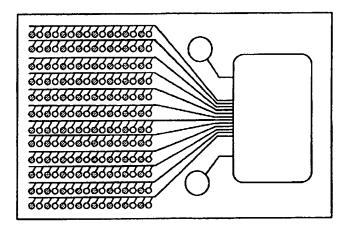


FIG. 9C

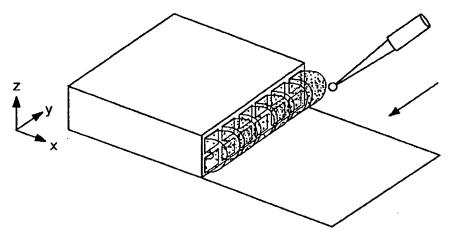


FIG. 10D

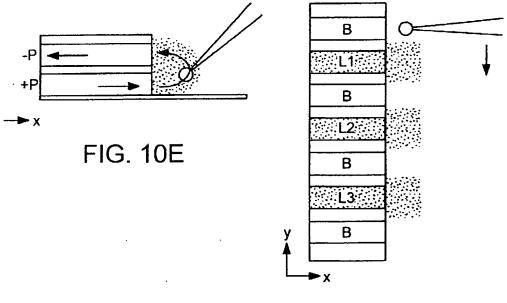
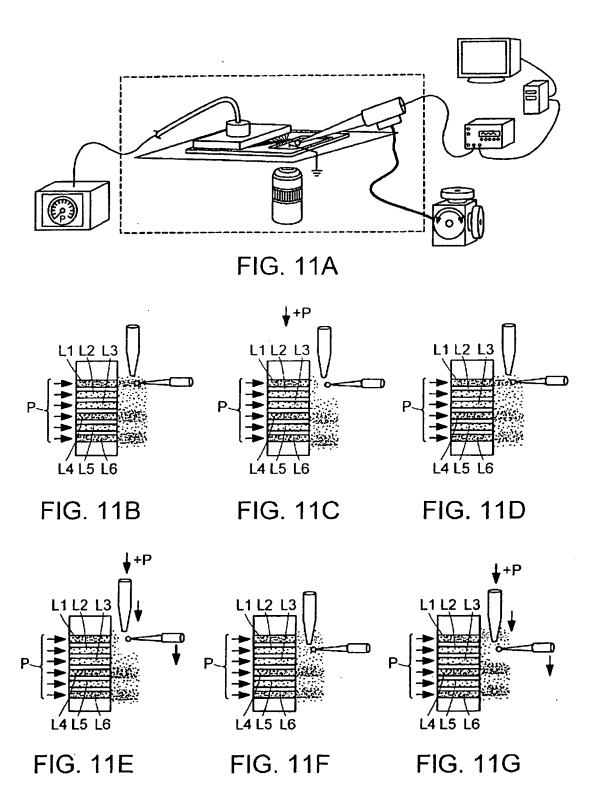
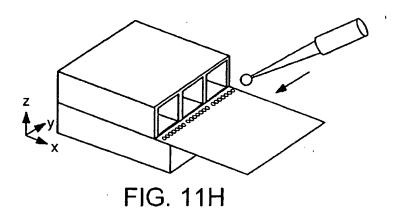
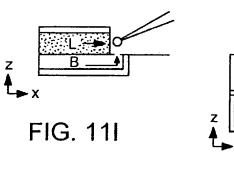
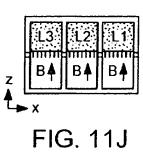


FIG. 10F









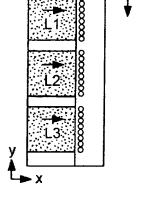


FIG. 11K

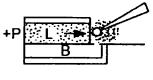


FIG. 11L

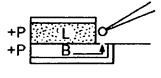


FIG. 11M

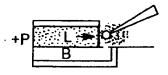
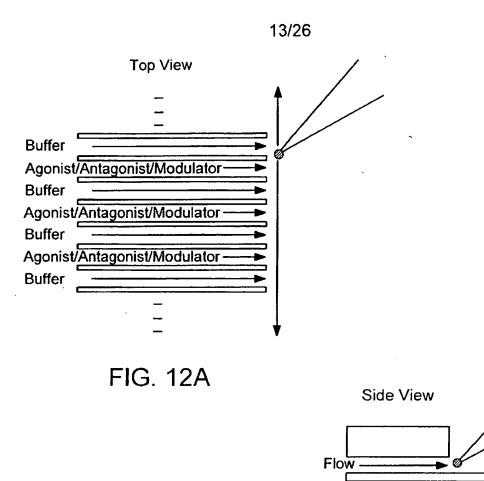
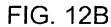
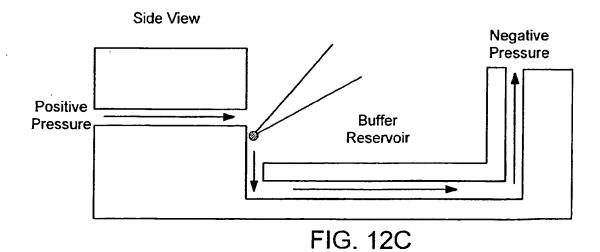


FIG. 11N





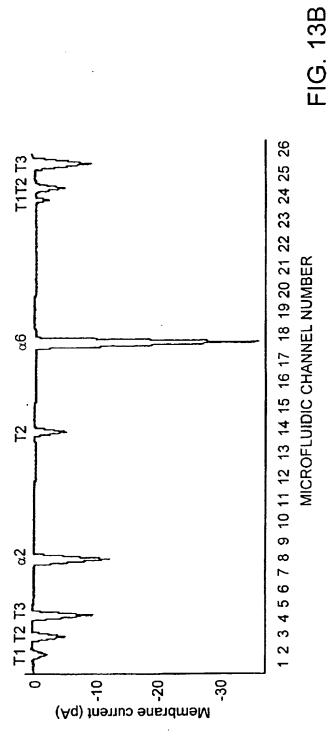


1	4/26

annel B T1 T2 ntent	nel 1 2 3
Т3	4
В	2
_	9
B α2	7 8
2 B	6
α3	
<u>в</u>	10 11 12 13 14
α 4	12
<u>B</u>	13
12	41
	15
α5	15 16 17 18 19
<u>a</u>	17
98	18
8	19
a7	20 21
ω	21
a.8	22
В	23
1	24
12	25
T3	26

FIG. 13A

Simulated trace for a single forward scan across microfluidic channel outlets:



Score sheet (mean peak current amplitude of 6 scans)

9	56					
5						
ļ	.25					
_	24					
0	23					
0	22					
0	21					
0	20					
0	19 20 21					
37	18					
0	17					
0	16					
0	15					
τ ₀	1 12 13 14 15 16 17 18					
0						
0						
0	11					
0	10 11					
0	6					
12	8					
0	2					
0	9					
0	5					
10	4					
5	က					
-	2					
0	_					
Receptor response	Channel #					

B=Buffer solution T1= Test compound with known efficacy (agonist) at low concentration

T2= Test compound with known efficacy (agonist) at medium concentration (close to EC_{50} -value)

T3= Test compound with known efficacy (agonist) at high concentration(saturating concentration).

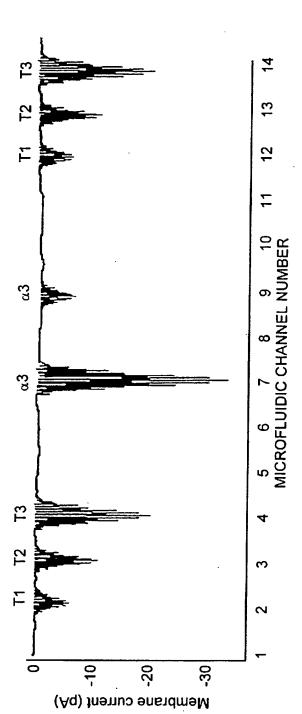
α=agonist with unknown efficacy

Channel #

Channel content

Simulated trace for a single forward scan across microfluidic channel outlets.

FIG. 14A



16/26

FIG. 14B

Score sheet (mean peak current amplitude of 6 scans)

20	14
10	13
2	12
0	1
0	10
4	6
0	æ
\$6	2
0	9
0	5
20	4
10	3
5	2
0	-
Receptor response	Channel #

B = Buffer solution T1 = Test compound with known efficacy (agonist) at low concentration

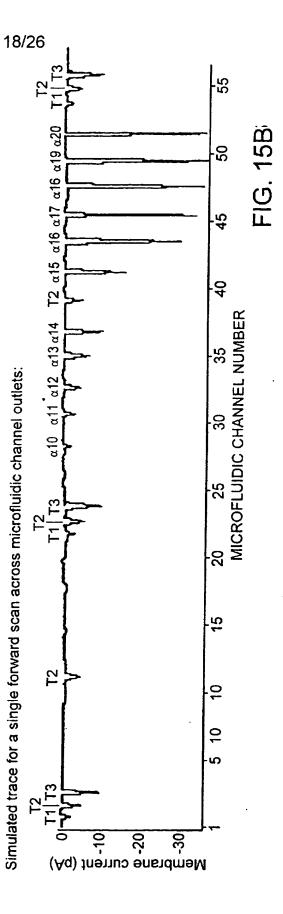
T2 = Test compound with known efficacy (agonist) at medium concentration (close to EC_{50} -value)

T3= Test compound with known efficacy (agonist) at high concentration(saturating concentration). α = agonist with unknown efficacy

FIG. 14C

α9	28	В	29
80	27	₽ €	
Т3	26	В	31 30
12	25	ಶ ∓	32
α8 B T1 T2 T3	24	മ	33 32
ω	23	2 Z	34
82	22	Ω.	35
ω	12 13 14 15 16 17 18 19 20 21 22 23	2 th	36
2α	20	α 1	43 42 41 40 39 38 37
В	19	8 4	38
σω	18	æ	39
80	17	72	40
ಶುಣ	16	മ	14
В	15	2 13	42
T2 B	14	8	43
В	13	α 16	44
g 4	12	Ω	45 44
ω	11	α 17	46
ರ ೧	9 10 11	В	47
В	တ	¤ 2 €	48 47
80	7 8	В	49
മ	^	2 G	50
۵1	မ	В	51
B	ည	Τ΄ α 3 20	52 51
⊢ ღ	4	— ო	53
T1 T2 T	1 2 3 4	T2	55 54 53
1	2	11	55
æ	1	മ	56
Content B in channel	Channel #	Content B T1 T2 T in channel	Channel #

FIG. 15A.



19/26

Score sheet (mean peak current amplitude of 6 scans)

Receptor	٥	-	ď	1	2 0 0	-	-		-	-	-		6	u		T-		-	H				 				-	
response	<u> </u>	-	,	2	,	,	>	>	>	>	>	>	>	2		 >		 >				 >	_			0	<u>-</u>	_
Channel #	1	2	2 3 4	4	5	9	2	ω	ი	10	=	12	13	9 10 11 12 13 14 15 16 17 18 19 20	15	19	17	18	6	ő	21 22	22	23 24		25	26 2	27 2	28
Receptor response	0	-	5 10 36	5		0	35	0 32 0 27 0 24	32	0	27	0	24	0 15 0 5	15	0	2	0 12 0 8	2	0	80	0	9	0	က	0 2		0
Channel #	56	55	54	53	52	51	50 49	49	48	47	46	45	44	45 44 43 42 41 40 39	42,	#	\$	1	38	37 36		35 34	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	33	32	33 32 31 30	9	29

B = Buffer solution

T1= Test compound with known efficacy (antagonist or agonist) at low concentration

T2= Test compound with known efficacy (antagonist or agonist) at medium concentration (close to EC₅₀-value)

T3= Test compound with known efficacy (antagonist or agonist) at high concentration(saturating concentration).

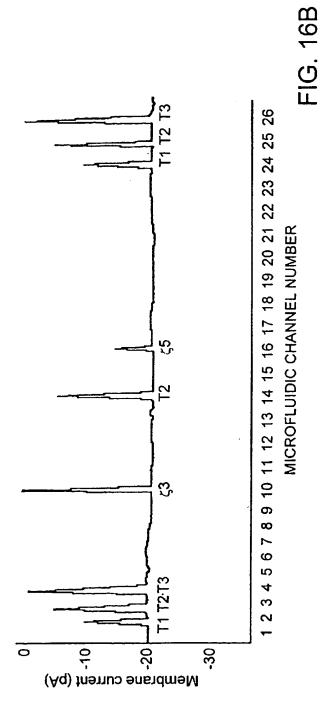
 α 1-to- α 28 agonist with unknown efficacy at different concentration progressively diluted(each step 10 times) to α 1

FIG. 15C

T3	26
12	25
T1 T2	24 25
	23
B: A B+ A+ B+ + +\zeta \ \ A \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \	22
₩ 4	19 20 21 22
みなと	20
œ + ∢	19
+ B + C + C + C + C + C + C + C + C + C	12 13 14 15 16 17 18
œ + ∢	17
Α [†] ν	16
∞ + ∢	15
72 B A + A + 45	4
m + ∢	13
4 [†] 4	12
α + α Α ⁺ , 4 α + α	11
Αţη	10 11
∞ + ∢	6
A 7, 2	8
യ + ∢	7
4 + +	9
B + A	5
+ \$ ∀	4
7 + A	က
⊢ ‡4	2
B + 4	-
Channel content	Channel #

FIG. 16A

Simulated trace for a single forward scan across microfluidic channel outlets:



Score sheet (mean peak current amplitude of 6 scans)

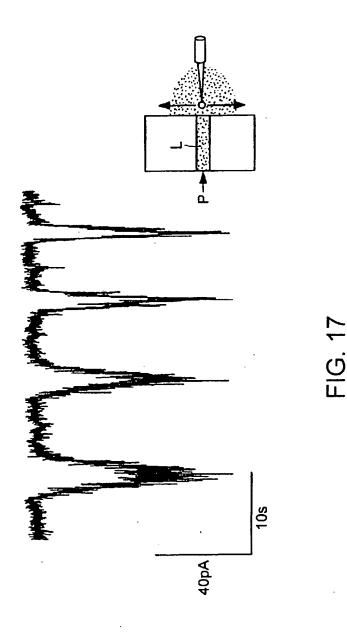
Receptor 20 response	20	10	5	-	20	20	20 ;	20 20	20		20	20	20	ည	20	15	20	 	50	20 20 5 20 15 20 20 20 20 20	20	20	20	10	ည	<u> </u>
Channel #	-	2	က	4	5	9	7	80	6	5	=	12	13	4	15	16	17	18	9	10 11 12 13 14 15 16 17 18 19 20 21 22	24		23	3 24	25	56

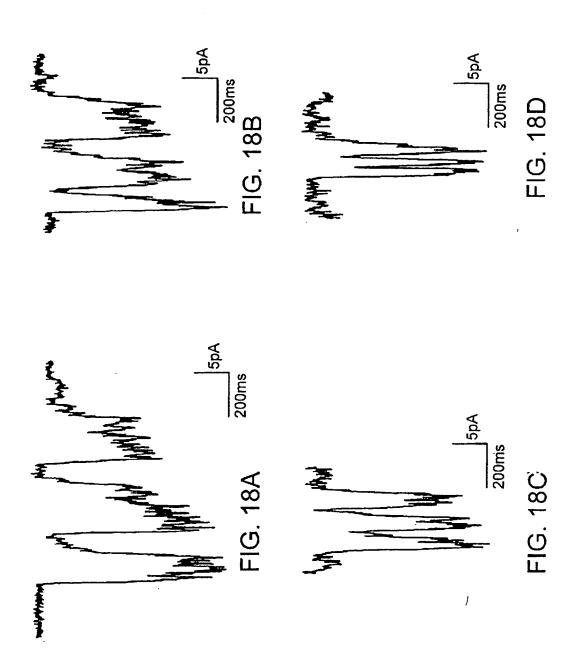
B=Buffer solution T1= Test compound with known efficacy (antagonist) at low concentration

T2= Test compound with known efficacy (antagonist) at medium concentration (close to EC_{50} -value)

T3= Test compound with known efficacy (antagonist) at high concentration(saturating concentration).

A= agonist with known efficacy ζ= antagonist with unknown efficacy





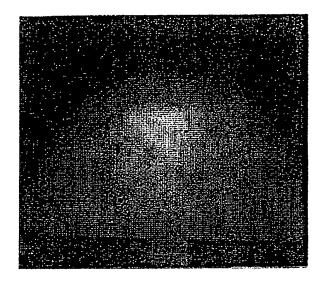


FIG. 19A

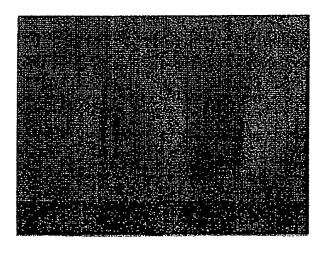


FIG. 19B



